

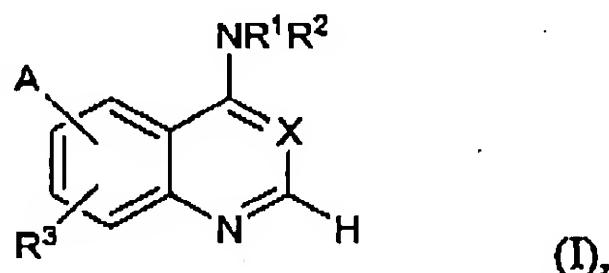
Application No.: 10/642,440

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AMENDMENT TO THE CLAIMS

A listing of the claims presented in this patent application appears below. This listing replaces all prior versions and listing of claims in this patent application.

Claim 1 (currently amended): A compound including resolved enantiomers, diastereomers[.,.] solvates and pharmaceutically acceptable salts thereof, said compound comprising Formula I:



wherein an A group is bonded to at least one of the carbons at the 5, 6, 7 or 8 position of the bicyclic ring, and the ring is substituted by up to three independent R³ groups;

X is N;

R¹ is a substituted or unsubstituted, monocyclic or bicyclic, aryl moiety;

R² is H or a substituted or unsubstituted C₁₋₈ alkyl[.,.]

or R² is a C₁₋₈ alkyl having a terminal carbon atom bound to one or the ring atoms of R¹;

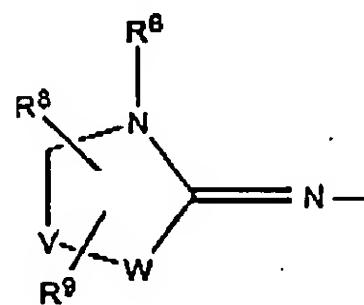
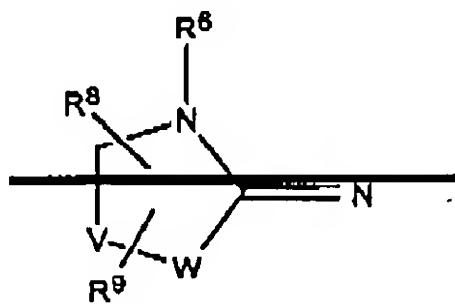
R³ is hydrogen, halogen, cyano, nitro, C₁-C₁₀ alkyl, C₂-C₁₀ alkenyl, C₂-C₁₀ alkynyl, C₃-C₁₀ cycloalkyl, C₃-C₁₀ cycloalkylalkyl, aryl, arylalkyl, heteroaryl, heteroarylalkyl, heterocyclyl, heterocyclylalkyl, -NR⁴SO₂R⁵, -SO₂NR⁶R⁴, -C(O)R⁶, -C(O)OR⁶, -OC(O)R⁶, -NR⁴C(O)OR⁵, -NR⁴C(O)R⁶, -C(O)NR⁴R⁶, -NR⁴R⁶, -NR⁴C(O)NR⁴R⁶, -OR⁶, -S(O)R⁵, -SO₂R⁵, where each of the above alkyl, alkenyl, alkynyl, cycloalkyl, aryl, heteroaryl and heterocyclyl portion of R³ is optionally substituted with one to five groups independently selected from oxo, halogen, cyano, nitro, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, -NR⁴SO₂R⁵, -SO₂NR⁶R⁴, -C(O)R⁶, -C(O)OR⁶, -OC(O)R⁶, -NR⁴C(O)OR⁵, -NR⁴C(O)CR⁶, -C(O)NR⁴R⁶, -NR⁴R⁶, -NR⁴C(O)NR⁴R⁶, -NR⁴C(NCN)NR⁴R⁶, -OR⁶, -S(O)R⁵, -SO₂R⁵, aryl, arylalkyl, heteroaryl, heteroarylalkyl, heterocyclyl, and heterocyclylalkyl;

A is -(U)_nZ, where

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n is 0 or 1, and U is C₁-C₄ alkyl, C₂-C₄ alkenyl or C₂-C₄ alkynyl; where each alkyl, alkenyl or alkynyl is optionally substituted with up to five groups independently selected from oxo, halogen, cyano, nitro, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, -NR⁴SO₂R⁵, -SO₂NR⁶R⁴, -C(O)R⁶, -C(O)OR⁶, -OC(O)R⁶, -NR⁴C(O)OR⁵, -NR⁴C(O)CR⁶, -C(O)NR⁴R⁶, -NR⁴R⁶, -NR⁴C(O)NR⁴R⁶, -NR⁴C(NCN)NR⁴R⁶, -OR⁶, -S(O)R⁵, -SO₂R⁵, aryl, arylalkyl, heteroaryl, heteroarylalkyl, heterocyclyl, and heterocyclylalkyl;

Z is



where W and V are selected independently from CR⁷R⁸, CR⁸R⁹, O, NR⁶, S, SO, SO₂, provided

if W is O, NR⁶, S, SO, SO₂, then V is CR⁸R⁹, and provided that NR⁶ of Z is not NH;

Z includes one or more R⁸ or R⁹ groups, wherein said R⁸ and R⁹ groups may be bonded to the same or different atoms;

R⁴ is H or C₁₋₆ alkyl;

R⁵ is trifluoromethyl, C₁-C₁₀ alkyl, C₃-C₁₀ cycloalkyl, aryl, arylalkyl, heteroaryl, heteroarylalkyl, heterocyclyl, heterocyclylalkyl, where each alkyl, cycloalkyl, aryl, heteroaryl, heterocyclyl and heterocyclylalkyl is optionally substituted with one to five groups independently selected from oxo, halogen, cyano, nitro, OR⁶, NR⁴R⁶, trifluoromethyl,

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difluoromethoxy, trifluoromethoxy, azido, aryl, heteroaryl, arylalkyl, heteroarylalkyl, heterocyclyl, and heterocyclalkyl;

R⁶, R⁸ and R⁹ are independently selected from hydrogen, trifluoromethyl, C₁-C₁₀ alkyl, (CH₂)₀₋₄C₃-C₁₀ cycloalkyl, aryl, arylalkyl, heteroaryl, heteroarylalkyl, heterocyclyl, heterocyclalkyl, where each alkyl, cycloalkyl, aryl, heteroaryl and heterocyclyl is optionally substituted with one to five groups independently selected from oxo, halogen, cyano, nitro, OR⁶, NR⁶R⁸, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, aryl, heteroaryl, arylalkyl, heteroarylalkyl, heterocyclyl, and heterocyclalkyl[[:]]:

R⁷ is hydrogen, halogen, cyano, nitro, C₁-C₁₀ alkyl, C₂-C₁₀ alkenyl, C₂-C₁₀ alkynyl, C₃-C₁₀ cycloalkyl, C₃-C₁₀ cycloalkylalkyl, aryl, arylalkyl, heteroaryl, heteroarylalkyl, heterocyclyl, heterocyclalkyl, -NR⁴SO₂R⁵, -SO₂NR⁶R⁴, -C(O)R⁶, -C(O)OR⁶, -OC(O)R⁶, -NR⁴C(O)OR⁵, -NR⁴C(O)R⁶, -C(O)NR⁴R⁶, -NR⁴R⁶, -NR⁴C(O)NR⁴R⁶, -OR⁶, -S(O)R⁵, -SO₂R⁵, where each of the above alkyl, alkenyl, alkynyl, cycloalkyl, aryl, heteroaryl and heterocyclyl portion of R³ is optionally substituted with one to five groups independently selected from oxo, halogen, cyano, nitro, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, -NR⁴SO₂R⁵, -SO₂NR⁶R⁴, -C(O)R⁶, -C(O)OR⁶, -OC(O)R⁶, -NR⁴C(O)OR⁵, -NR⁴C(O)CR⁶, -C(O)NR⁴R⁶, -NR⁴R⁶, -NR⁴C(O)NR⁴R⁶, -NR⁴C(NCN)NR⁴R⁶, -OR⁶, -S(O)R⁵, -SO₂R⁵, aryl, arylalkyl, heteroaryl, heteroarylalkyl, heterocyclyl, and heterocyclalkyl;

an R⁴ group and an R⁶ group may be independently joined to complete a 3 to 10 membered cyclic ring optionally containing additional heteroatoms selected from the group consisting of O, S, SO, SO₂ and NR⁶ where each ring carbon may be optionally substituted with one to three groups independently selected from halogen, cyano, nitro, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, aryl, OR⁸, NR⁶R⁸, heteroaryl, arylalkyl, heteroarylalkyl, heterocyclyl, and heterocyclalkyl; provided said ring does not contain two adjacent O or two adjacent S atoms;

an R⁶ group and an R⁸ group may be independently joined to complete a 3 to 10 membered cyclic ring optionally containing additional heteroatoms selected from the group consisting of O, S, SO, SO₂ and NR⁶ where each ring carbon may be optionally substituted with one to three groups independently selected from halogen, cyano, nitro, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, aryl, OR⁸, NR⁶R⁸, heteroaryl, arylalkyl,

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heteroarylalkyl, heterocyclyl, and heterocyclylalkyl; provided said ring does not contain two adjacent O or two adjacent S atoms;

an R⁷ group and an R⁸ group may be independently joined to complete a 3 to 10 membered cyclic ring optionally containing additional heteroatoms selected from the group consisting of O, S, SO, SO₂ and NR⁶ where each ring carbon may be optionally substituted with one to three groups independently selected from halogen, cyano, nitro, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, aryl, OR⁸, NR⁶R⁸, heteroaryl, arylalkyl, heteroarylalkyl, heterocyclyl, and heterocyclylalkyl; provided said ring does not contain two adjacent O or two adjacent S atoms; and

an R⁸ group and an R⁹ group may be independently joined to complete a 3 to 10 membered cyclic ring optionally containing additional heteroatoms selected from the group consisting of O, S, SO, SO₂ and NR⁶ where each ring carbon may be optionally substituted with one to three groups independently selected from halogen, cyano, nitro, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, aryl, OR⁸, NR⁶R⁸, heteroaryl, arylalkyl, heteroarylalkyl, heterocyclyl, and heterocyclylalkyl; provided said ring does not contain two adjacent O or two adjacent S atoms.

Claim 2 (original): The compound of claim 1, wherein R² is a C₁₋₈ alkyl having a terminal carbon atom bound to one of the ring atoms of R¹.

Claim 3 (original): The compound of claim 1, wherein an A group is bonded to at least one of the carbons at the 6 or 7 position of the bicyclic ring.

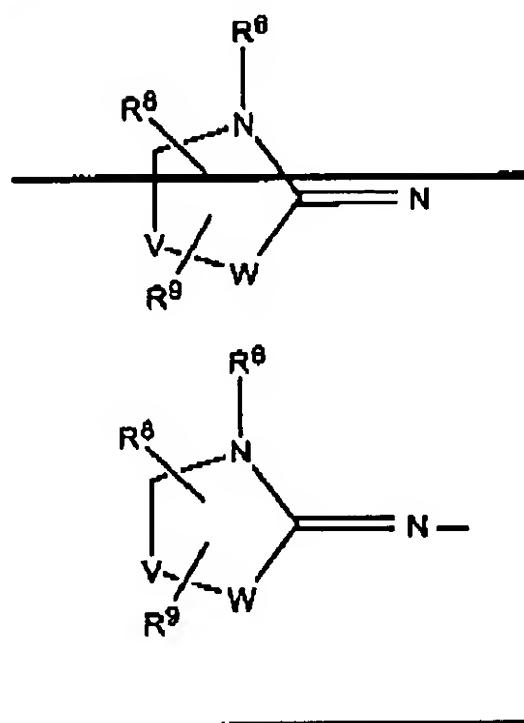
Claim 4 (previously amended): The compound of claim 1, wherein R² is hydrogen, and R³ is hydrogen or OR⁶.

Claim 5 (previously amended): The compound of claim 3, wherein R³ is hydrogen or OR⁶, and n is 0.

Claim 6 (original): The compound of claim 1, wherein R² is hydrogen.

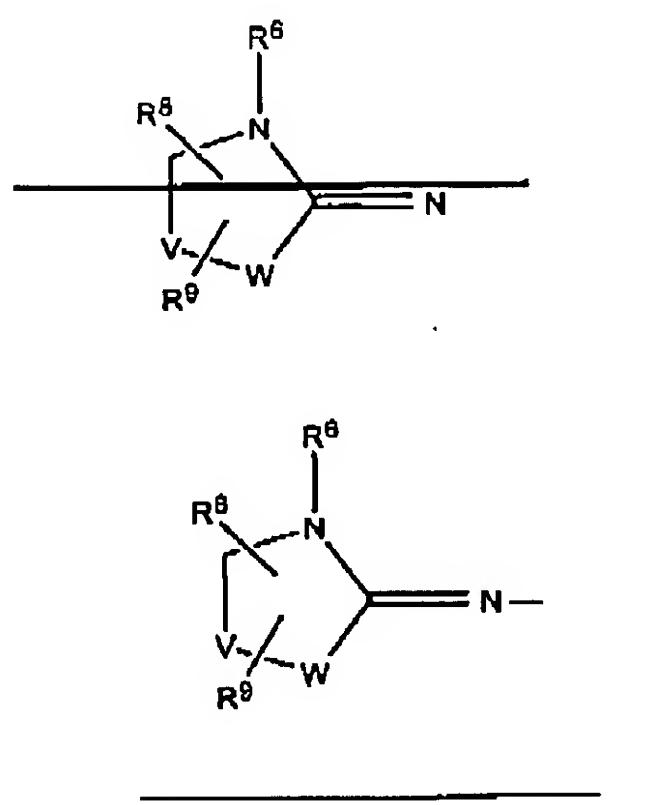
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Claim 7 (currently amended): The compound of claim 1, wherein Z is



and W is O.

Claim 8 (currently amended): The compound of claim 5, wherein Z is



and W is O.

Claim 9 (original): The compound of claim 1, wherein the R⁴ group and the R⁶ group are independently joined to complete a 3 to 10 membered cyclic ring optionally containing

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additional heteroatoms selected from the group consisting of O, S, SO, SO₂ and NR⁶ where each ring carbon may be optionally substituted with one to three groups independently selected from halogen, cyano, nitro, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, aryl, OR⁸, NR⁶R⁸, heteroaryl, arylalkyl, heteroarylalkyl, heterocyclyl, and heterocyclylalkyl; provided said ring does not contain two adjacent O or two adjacent S atoms.

Claim 10 (original): The compound of claim 1, wherein the R⁶ group and the R⁸ group are independently joined to complete a 3 to 10 membered cyclic ring optionally containing additional heteroatoms selected from the group consisting of O, S, SO, SO₂ and NR⁶ where each ring carbon may be optionally substituted with one to three groups independently selected from halogen, cyano, nitro, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, aryl, OR⁸, NR⁶R⁸, heteroaryl, arylalkyl, heteroarylalkyl, heterocyclyl, and heterocyclylalkyl; provided said ring does not contain two adjacent O or two adjacent S atoms.

Claim 11 (previously amended): The compound of claim 1, wherein the R⁷ group and the R⁸ group are independently joined to complete a 3 to 10 membered cyclic ring optionally containing additional heteroatoms selected from the group consisting of O, S, SO, SO₂ and NR⁶ where each ring carbon may be optionally substituted with one to three groups independently selected from halogen, cyano, nitro, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, aryl, OR⁸, NR⁶R⁸, heteroaryl, arylalkyl, heteroarylalkyl, heterocyclyl, and heterocyclylalkyl; provided said ring does not contain two adjacent O or two adjacent S atoms.

Claim 12 (original): The compound of claim 1, wherein the R⁸ group and the R⁹ group are independently joined to complete a 3 to 10 membered cyclic ring optionally containing additional heteroatoms selected from the group consisting of O, S, SO, SO₂ and NR⁶ where each ring carbon may be optionally substituted with one to three groups independently selected from halogen, cyano, nitro, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, aryl, OR⁸, NR⁶R⁸, heteroaryl, arylalkyl, heteroarylalkyl, heterocyclyl, and heterocyclylalkyl; provided said ring does not contain two adjacent O or two adjacent S atoms.

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Claim 13 (withdrawn): A method of treating hyperproliferative diseases in a mammal comprising administering a therapeutically effective amount of the compound defined in claim 1 to said mammal.

Claim 14 (withdrawn): A method of treating hyperproliferative diseases in a mammal comprising administering a therapeutically effective amount of the compound defined in claim 2 to said mammal.

Claim 15 (withdrawn): A method of treating hyperproliferative diseases in a mammal comprising administering a therapeutically effective amount of the compound defined in claim 3 to said mammal.

Claim 16 (withdrawn): A method of treating hyperproliferative diseases in a mammal comprising administering a therapeutically effective amount of the compound defined in claim 4 to said mammal.

Claim 17 (withdrawn): A method of treating hyperproliferative diseases in a mammal comprising administering a therapeutically effective amount of the compound defined in claim 5 to said mammal.

Claim 18 (withdrawn): A method of treating hyperproliferative diseases in a mammal comprising administering a therapeutically effective amount of the compound defined in claim 6 to said mammal.

Claim 19 (withdrawn): A method of treating hyperproliferative diseases in a mammal comprising administering a therapeutically effective amount of the compound defined in claim 7 to said mammal.

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Claim 20 (withdrawn): A method of treating hyperproliferative diseases in a mammal comprising administering a therapeutically effective amount of the compound defined in claim 8 to said mammal.

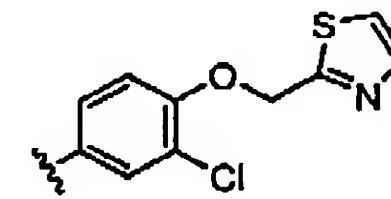
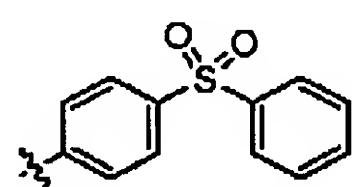
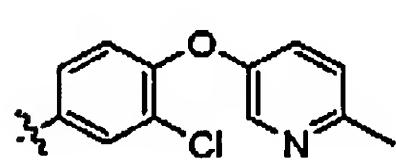
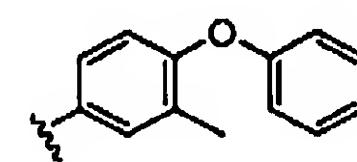
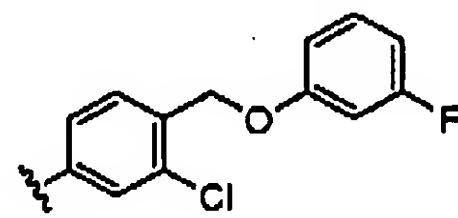
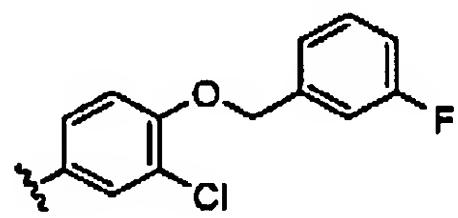
Claim 21 (withdrawn): A method of treating hyperproliferative diseases in a mammal comprising administering a therapeutically effective amount of the compound defined in claim 9 to said mammal.

Claim 22 (withdrawn): A method of treating hyperproliferative diseases in a mammal comprising administering a therapeutically effective amount of the compound defined in claim 10 to said mammal.

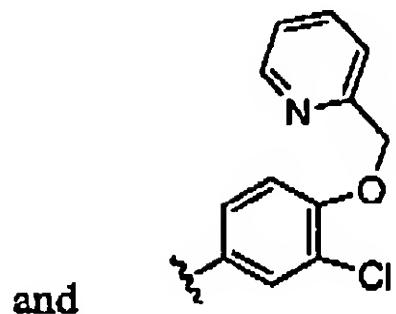
Claim 23 (withdrawn): A method of treating hyperproliferative diseases in a mammal comprising administering a therapeutically effective amount of the compound defined in claim 11 to said mammal.

Claim 24 (withdrawn): A method of treating hyperproliferative diseases in a mammal comprising administering a therapeutically effective amount of the compound defined in claim 12 to said mammal.

Claim 25 (previously presented): The compound of claim 1, wherein R¹ is selected from the structures:



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Claim 26 (previously presented): The compound of claim 7, wherein R⁶ is an optionally substituted alkyl or cycloalkyl.

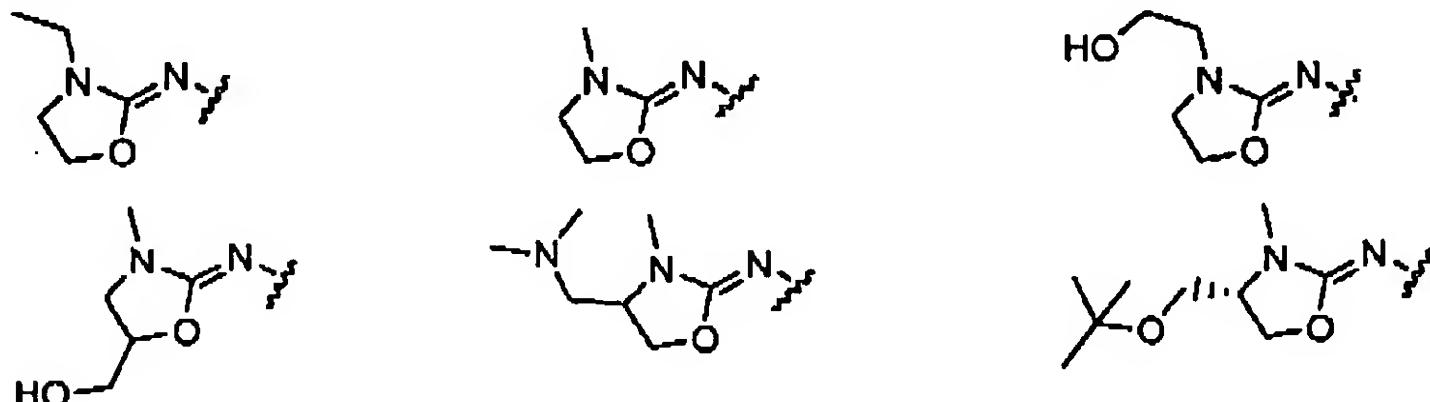
Claim 27 (previously presented): The compound of claim 26, wherein R⁶ is methyl, ethyl, CH₂CF₃, CH₂CH₂OH, or cyclopropyl.

Claim 28 (previously presented): The compound of claim 26, wherein R⁸ and R⁹ are independently an optionally substituted alkyl.

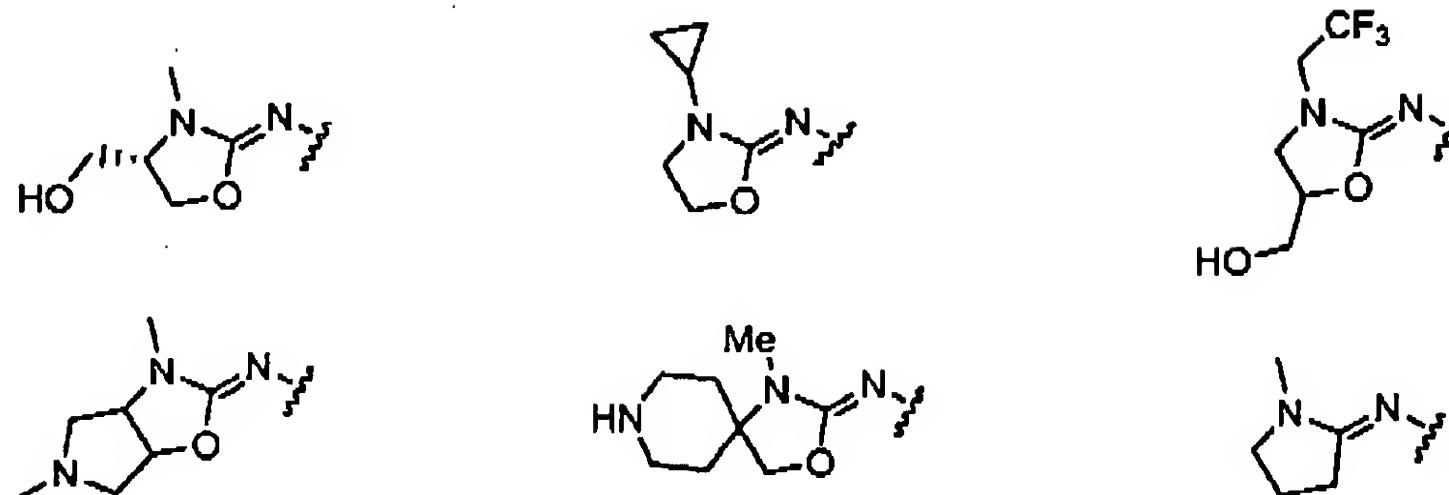
Claim 29 (previously presented): The compound of claim 28, wherein R⁸ and R⁹ are independently CH₂OH, CH₂NMe₂ or CH₂O-t-butyl.

Claim 30 (previously presented): The compound of claim 26, wherein R⁸ and R⁹ together with the atoms to which they are attached form an optionally substituted heterocyclic ring.

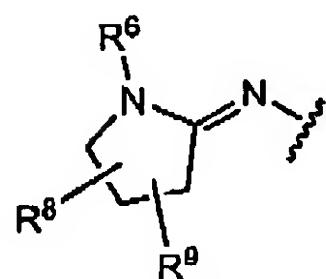
Claim 31 (currently amended): The compound of claim 7, wherein Z is selected from the structures:



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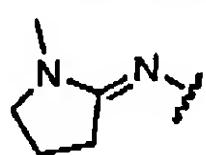
Claim 32 (previously presented): The compound of claim 1, wherein Z is



Claim 33 (previously presented): The compound of claim 32, wherein R⁶ is an optionally substituted alkyl.

Claim 34 (previously amended): The compound of claim 33, wherein Z is methyl.

Claim 35 (previously presented): The compound of claim 34, wherein Z is



Claim 36 (currently amended): The compound of claim 1, selected from:

N4-[3-Chloro-4-(3-fluorobenzylxy)-phenyl]-N6-(3-methyl-oxazolidin-2-ylidene)-quinazoline-4,6-diamine;

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N-4-[3-Chloro-4-(3-fluorobenzylxy)-phenyl]-N6-(3-ethyl-oxazolidin-2-ylidene)-quinazoline-4,6-diamine;

(2-{4-[3-Chloro-4-(3-fluorobenzylxy)-phenylamino]-quinazolin-6-ylimino}-3-methyl-oxazolidin-5-yl)-methanol;

2-(2-{4-[3-Chloro-4-(3-fluorobenzylxy)-phenylamino]-quinazolin-6-ylimino}-oxazolidin-3-yl)-ethanol;

N-4-[3-Chloro-4-(3-fluorobenzylxy)-phenyl]-N6-(4-dimethylaminomethyl-3-methyl-oxazolidin-2-ylidene)-quinazoline-4,6-diamine;

(S)-N6-(4-tert-Butoxymethyl-3-methyl-oxazolidin-2-ylidene)-N4-[3-chloro-4-(3-fluorophenoxyethyl)-phenyl]-quinazoline-4,6-diamine;

(S)-(2-{4-[3-Chloro-4-(3-fluorophenoxyethyl)-phenylamino]-quinazolin-6-ylimino}-3-methyl-oxazolidin-5-yl)-methanol;

(2-{4-[3-Chloro-4-(3-fluorophenoxyethyl)-phenylamino]-quinazolin-6-ylimino}-3-methyl-oxazolidin-5-yl)-methanol;

(3-Methyl-2-[4-(3-methyl-4-phenoxyphenylamino)-quinazolin-6-ylimino]-oxazolidin-5-yl)-methanol;

(2-{4-[3-Chloro-4-(6-methylpyridin-3-yloxy)-phenylamino]-quinazolin-6-ylimino}-3-methyl-oxazolidin-5-yl)-methanol;

N4-(4-Benzenesulfonylphenyl)-N6-(3-methyloxazolidin-2-ylidene)-quinazoline-4,6-diamine;

{2-[4-(4-Benzenesulfonylphenylamino)-quinazolin-6-ylimino]-3-methyl-oxazolidin-5-yl}-methanol;

N4-(4-Benzenesulfonylphenyl)-N6-(3-cyclopropyloxazolidin-2-ylidene)-quinazoline-4,6-diamine;

N6-(Dimethylhexahydropyrrolo[3,4-d]oxazol-2-ylidene)-N4-(3-methyl-4-phenoxyphenyl)-quinazoline-4,6-diamine;

N4-[3-Chloro-4-(thiazol-2-ylmethoxy)-phenyl]-N6-(3-methyloxazolidin-2-ylidene)-quinazoline-4,6-diamine;

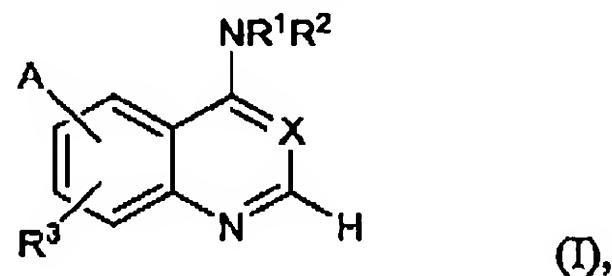
N4-[3-Chloro-4-(pyridin-2-ylmethoxy)-phenyl]-N6-(dimethyl-3-oxa-1,8-diaza-spiro[4.5]dec-2-ylidene)-quinazoline-4,6-diamine;

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[2-{4-[3-Chloro-4-(3-fluorobenzyl)oxy]-phenylamino}-quinazolin-6-ylimino]-3-(2,2,2-trifluoroethyl)-oxazolidin-5-yl]-methanol; and

N4-[3-Chloro-4-(3-fluorobenzyl)oxy]-phenyl]-N6-(1-methylpyrrolidin-2-ylidene)-quinazoline-4,6-diamine.

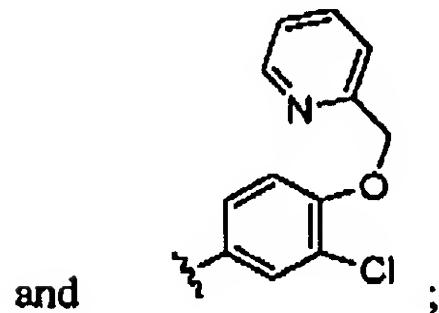
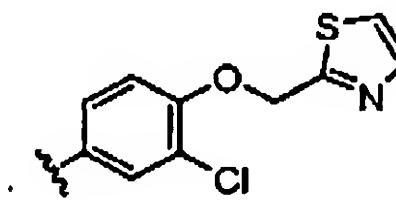
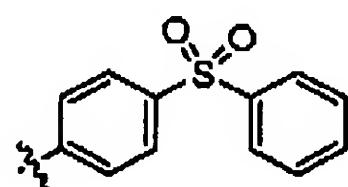
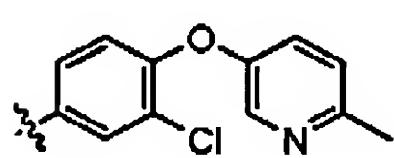
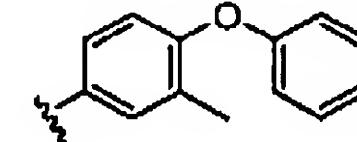
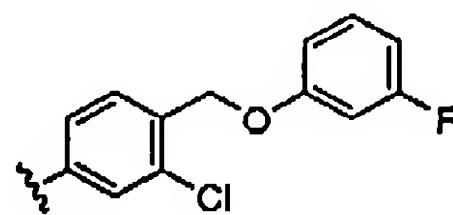
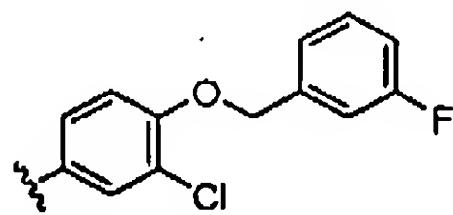
Claim 37 (new): A compound including resolved enantiomers, diastereomers and pharmaceutically acceptable salts thereof, said compound comprising Formula I:



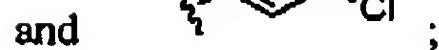
wherein

X is N;

R¹ is selected from the structures:



and



R² is hydrogen or a substituted or unsubstituted C₁₋₈ alkyl;

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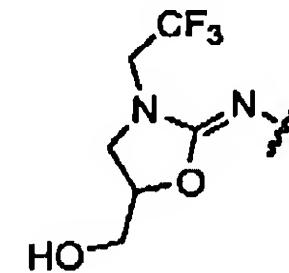
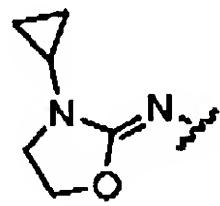
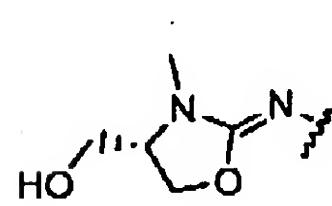
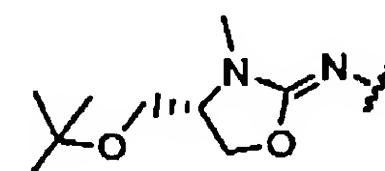
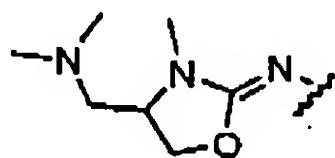
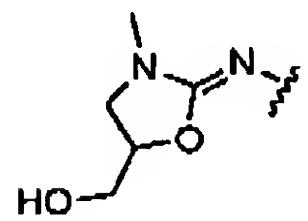
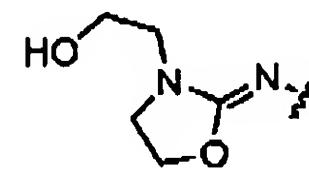
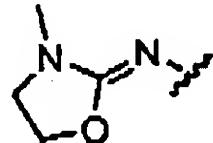
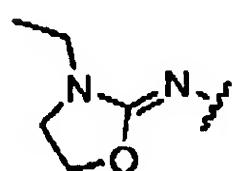
R^3 is hydrogen, halogen, cyano, nitro, C_1-C_{10} alkyl, C_2-C_{10} alkenyl, C_2-C_{10} alkynyl, C_3-C_{10} cycloalkyl, C_3-C_{10} cycloalkylalkyl, aryl, arylalkyl, heteroaryl, heteroarylalkyl, heterocyclyl, heterocyclylalkyl, $-NR^4SO_2R^5$, $-SO_2NR^6R^4$, $-C(O)R^6$, $-C(O)OR^6$, $-OC(O)R^6$, $-NR^4C(O)OR^5$, $-NR^4C(O)R^6$, $-C(O)NR^4R^6$, $-NR^4R^6$, $-NR^4C(O)NR^4R^6$, $-OR^6$, $-S(O)R^5$, $-SO_2R^5$, where each of the above alkyl, alkenyl, alkynyl, cycloalkyl, aryl, heteroaryl and heterocyclyl portion of R^3 is optionally substituted with one to five groups independently selected from oxo, halogen, cyano, nitro, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, $-NR^4SO_2R^5$, $-SO_2NR^6R^4$, $-C(O)R^6$, $-C(O)OR^6$, $-OC(O)R^6$, $-NR^4C(O)OR^5$, $-NR^4C(O)CR^6$, $-C(O)NR^4R^6$, $-NR^4R^6$, $-NR^4C(O)NR^4R^6$, $-NR^4C(NCN)NR^4R^6$, $-OR^6$, $-S(O)R^5$, $-SO_2R^5$, aryl, arylalkyl, heteroaryl, heteroarylalkyl, heterocyclyl, and heterocyclylalkyl;

A is $-(U)_nZ$, where

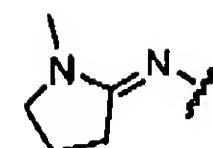
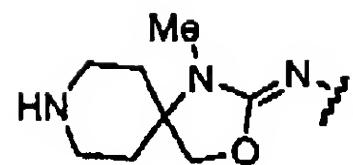
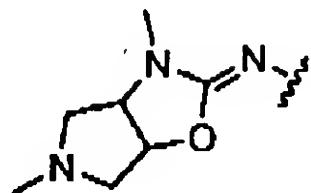
n is 0 or 1;

U is C_1-C_4 alkyl, C_2-C_4 alkenyl or C_2-C_4 alkynyl; where each alkyl, alkenyl or alkynyl is optionally substituted with up to five groups independently selected from oxo, halogen, cyano, nitro, trifluoromethyl, difluoromethoxy, trifluoromethoxy, azido, $-NR^4SO_2R^5$, $-SO_2NR^6R^4$, $-C(O)R^6$, $-C(O)OR^6$, $-OC(O)R^6$, $-NR^4C(O)OR^5$, $-NR^4C(O)CR^6$, $-C(O)NR^4R^6$, $-NR^4R^6$, $-NR^4C(O)NR^4R^6$, $-NR^4C(NCN)NR^4R^6$, $-OR^6$, $-S(O)R^5$, $-SO_2R^5$, aryl, arylalkyl, heteroaryl, heteroarylalkyl, heterocyclyl, and heterocyclylalkyl; and

Z is selected from the following structures:



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Claim 38 (new): The compound of claim 37, wherein n=0.

Claim 39 (new): The compound of claim 38, wherein R² is hydrogen.

Claim 40 (new): The compound of claim 38, wherein R³ is hydrogen.